

Application No.: 10/081,642

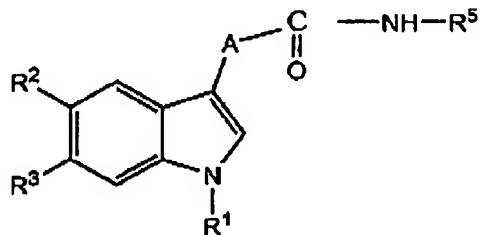
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IN THE CLAIMS

Claims 1-63 (canceled)

64. (new) A process for preparing a compound of formula 1



or a salt thereof, wherein

R¹ is a straight or branched C₁₋₁₂ alkyl optionally substituted with phenyl, or C₃₋₈ cycloalkyl radical wherein the phenyl radical is optionally substituted with a halo, nitro, hydroxy, C₁₋₄ alkyl, C₁₋₄ alkoxy, or COOH;

R² and R³ are each independently of each other hydrogen or an OH radical where at least one of R² and R³ are —OH;

R⁵ is a phenyl or pyridyl radical substituted with at least one halogen radical and is optionally further substituted with -H, -OH, -SH, -NH₂, -NHC₁₋₆ alkyl, -N(C₁₋₆ alkyl)₂, -NHC₆₋₁₄ aryl, -N(C₆₋₁₄ aryl)₂, -N(C₁₋₆ alkyl)(C₆₋₁₄ aryl), -NHCOR⁶, -NO₂, -CN, -COOH, -(CO)R⁶, -(CS)R⁶, -F, -Cl, -Br, -I, -O-C₁₋₆ alkyl, -O-C₆₋₁₄ aryl, -O(CO)R⁶, -S-C₁₋₆ alkyl, -S-C₆₋₁₄ aryl, -SOR⁶, or -SO₂R⁶; and

A is a bond, C=O, or a CHOH radical or a pharmaceutically acceptable salt thereof,

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which method comprises converting a compound of formula (I), wherein R² or R³ or R² and R³ are O-R⁷, into the compound of formula (I) by removing R⁷, wherein R⁷ is a substituent that is a protecting group selected from the group consisting of alkyl and aralkyl Lewis acid to cleave the ether and remove R⁷, to yield the compound of formula (I), wherein said compound of formula (I) is selected from the group consisting of N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2- hydroxyacetamide, N-(3,5-dichloropyridin-4-yl)-2-(1-(2,6-difluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide, N-(3,5-dichloropyridin-4-yl)-2-(1-(3-nitrobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide, N-(3,5-dichloropyridin-4-yl)-2-(1-propyl-5-hydroxyindol-3-yl)-2-oxoacetamide, N-(3,5-dichloropyridin-4-yl)-2-(1-isopropyl-5-hydroxyindol-3-yl)-2-oxoacetamide, N-(3,5-dichloropyridin-4-yl)-2-(1-cyclopentylmethyl-5-hydroxyindol-3-yl)-2 -oxoacetamide, N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-6-hydroxyindol-3-yl)-2- oxoacetamide, N-(3,5-dichloropyridin-4-yl)-5-hydroxy-1-(4-methoxybenzyl)indole-3-carboxamide and N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2- oxoacetamide.

65. (new) The method of claim 64, wherein R⁵ is substituted with one or two halogens.

66. (new) The method of claim 64, wherein R¹ is an optionally substituted C₁-C₂ alkyl.

67. (new) The method of claim 66, wherein R¹ is an optionally substituted C₁-C₂ alkyl.

68. (new) The method of claim 64, wherein R⁷ is methyl or ethyl.

69. (new) The method of claim 68, wherein R⁷ is methyl.

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70. (new) The method of claim 64, wherein the Lewis acid is selected from the group consisting of BBr_3 and $AlCl_3$.

71. (new) The method of claim 70, wherein said Lewis acid is BBr_3 .

72. (new) The method of claim 70, wherein removal of R^7 is in the presence of an additional activator.

73. (new) The method of claim 72, wherein the additional activator is selected from the group consisting of ethane-1,2-dithiol and benzyl mercaptan.

74. (new) The method of claim 70, wherein said Lewis acid is $AlCl_3$.

75. (new) The method of claim 64, wherein the ether cleavage is conducted at elevated or normal pressure.

76. (new) The method of claim 64, wherein the ether cleavage takes place in the presence of a suitable catalyst.

77. (new) A method for producing N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide by reacting a solution of N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-methoxyindol-3-yl)-2-oxoacetamide with BBr_3 while heating to form a heated solution, cooling the heated solution to yield a cooled solution, and mixing the cooled solution with an aqueous sodium hydrogencarbonate solution to crystallize the N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide.

78. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide sodium salt.

79. (new) The method of claim 77, wherein the solution is stirred during heating.

80. (new) The method of claim 79, wherein the solution is stirred during cooling.

81. (new) The method of claim 77, further comprising recovering the crystallized N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide.

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82. (new) The method of claim 80, further comprising recovering the crystallized N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide.

83. (new) The method of claim 82, wherein the solution is cooled to 20°C.

84. (new) The method of claim 77, further comprising recrystallizing the crystallized N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide.

85. (new) The method of claim 64, wherein the compound is a pharmaceutically acceptable salt of the compound.

86. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-5-hydroxyindol-3-yl)-2-hydroxyacetamide.

87. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(-(2,6-difluorobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide.

88. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(1-(3-nitrobenzyl)-5-hydroxyindol-3-yl)-2-oxoacetamide.

89. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(1-propyl-5-hydroxyindol-3-yl)-2-oxoacetamide.

90. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(1-isopropyl-5-hydroxyindol-3-yl)-2-oxoacetamide.

91. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(1-cyclopentylmethyl-5-hydroxyindol-3-yl)-2-oxoacetamide.

92. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-2-(1-(4-fluorobenzyl)-6-hydroxyindol-3-yl)-2-oxoacetamide.

93. (new) The method of claim 64, wherein the compound is N-(3,5-dichloropyridin-4-yl)-5-hydroxy-1-(4-methoxybenzyl)indole-3-carboxamide.